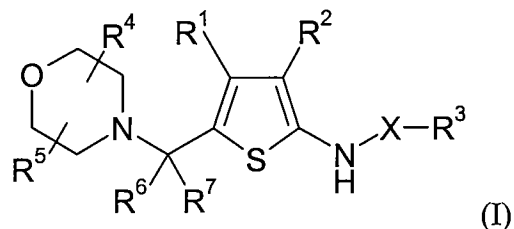


CLAIM AMENDMENTS

1. (original) A compound of the formula (I):



or a salt, solvate or N-oxide thereof, wherein:

R¹ and R² are the same or different and each is selected from hydrogen, saturated C₁₋₃ hydrocarbyl, halogen and cyano;

X is selected from C=O, C=S, C(=O)NH, C(=S)NH, C(=O)O, C(=O)S, C(=S)O and C(=S)S;

R³ is selected from aryl and heteroaryl groups each having from 5 to 12 ring members and being unsubstituted or substituted by one or more substituent groups R¹⁰;

R¹⁰ is selected from halogen, hydroxy, trifluoromethyl, cyano, nitro, carboxy, amino, mono- or di-C₁₋₄ hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members; a group R^a-R^b wherein R^a is a bond, O, CO, X¹C(X²), C(X²)X¹, X¹C(X²)X¹, S, SO, SO₂, NR^c, SO₂NR^c or NR^cSO₂; and R^b is selected from hydrogen, carbocyclic and heterocyclic groups having from 3 to 12 ring members, and a C₁₋₈ hydrocarbyl group optionally substituted by one or more substituents selected from hydroxy, oxo, halogen, cyano, nitro, carboxy, amino, mono- or di-C₁₋₄ hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members and wherein one or more carbon atoms of the C₁₋₈ hydrocarbyl group may optionally be replaced by O, S, SO, SO₂, NR^c, X¹C(X²), C(X²)X¹ or X¹C(X²)X¹; or two adjacent groups R¹⁰, together with the carbon atoms or heteroatoms to which they are attached may form a 5-membered heteroaryl ring or a 5- or 6-membered non-aromatic heterocyclic ring, wherein the said heteroaryl and heterocyclic groups contain up to 3 heteroatom ring members selected from N, O and S;

R^c is selected from hydrogen and C₁₋₄ hydrocarbyl; and

X^1 is O, S or NR^c and X^2 is $=O$, $=S$ or $=NR^c$;

R^4 and R^5 are the same or different and are selected from hydrogen and methyl; or one of R^4 and R^5 is selected from hydroxymethyl and ethyl and the other is hydrogen; and

R^6 and R^7 are the same or different and are selected from hydrogen and methyl.

2. (original) A compound according to claim 1 wherein R^3 is a monocyclic aryl or heteroaryl group.

3-82 (canceled)

83. (new) A compound according to claim 2 wherein the aryl group or heteroaryl group R^3 contains one or more substituent groups R^{10} selected from halogen, carbocyclic and heterocyclic groups having from 4 to 7 ring members and optionally substituted C_{1-8} hydrocarbyl groups.

84. (new) A compound according to claim 83 wherein the group R^3 contains a substituent R^{10} which is a carbocyclic or heterocyclic group having from 4 to 7 ring members and said carbocyclic or heterocyclic group is linked to the aryl or heteroaryl ring via a carbon nitrogen bond.

85. (new) A compound according to claim 84 wherein the carbocyclic or heterocyclic group R^{10} is a 4 to 7 membered heterocyclic group R^8 selected from morpholine, piperidino, piperazino, N-methyl piperazino, tetrahydrofuranyl and pyrrolidino.

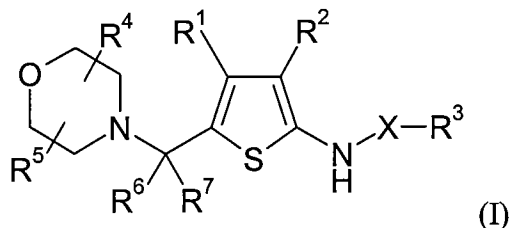
86. (new) A compound according to claim 1 wherein X is $C=O$ or $C(=O)NH$.

87. (new) A compound according to claim 1 wherein R^1 is selected from hydrogen, saturated C_{1-3} hydrocarbyl and halogen.

88. (new) A compound according to claim 1 wherein R^2 is selected from hydrogen, saturated C_{1-3} hydrocarbyl and halogen.

89. (new) A compound according to claim 87 wherein R^1 is chlorine.

90. (new) A compound according to claim 88 wherein R^2 is methyl.
91. (new) A compound according to claim 1 wherein R^4 and R^5 are both hydrogen.
92. (new) A compound according to claim 1 wherein R^6 and R^7 are both hydrogen.
93. (new) A compound according to claim 1 which is selected from:
N-(4-chloro-3-methyl-5-(morpholin-yl methyl-thiophen-2-yl)-3-fluoro-morpholin-4-yl-benzamide;
1-[5-tert-butyl-2(4-fluoro-phenyl)-2H-pyrazol-3-yl]-3-(4-chloro-3-methyl-5-morpholin-4-ylmethyl-thiophen-2-yl) urea;
1-[5-tert-butyl-2-(2,4-difluoro-phenyl)-2H-pyrazol-3-yl]-3-(4-chloro-3-methyl-5-morpholin-4-ylmethyl-thiophen-2-yl)-urea; and
1-(4-chloro-3-methyl-5-morpholin-4-ylmethyl-thiophen-2-yl)-3-[5-(tetrahydro-furan-2-yl)-[1,3,4]thiadiazol-2-yl]-urea.
94. (new) A pharmaceutical composition comprising a compound of the formula (I):



or a salt, solvate or N-oxide thereof, wherein:

R^1 and R^2 are the same or different and each is selected from hydrogen, saturated C_{1-3} hydrocarbyl, halogen and cyano;

X is selected from C=O, C=S, C(=O)NH, C(=S)NH, C(=O)O, C(=O)S, C(=S)O and C(=S)S;

R^3 is selected from aryl and heteroaryl groups each having from 5 to 12 ring members and being unsubstituted or substituted by one or more substituent groups R^{10} ;

R^{10} is selected from halogen, hydroxy, trifluoromethyl, cyano, nitro, carboxy,

amino, mono- or di-C₁₋₄ hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members; a group R^a-R^b wherein R^a is a bond, O, CO, X¹C(X²), C(X²)X¹, X¹C(X²)X¹, S, SO, SO₂, NR^c, SO₂NR^c or NR^cSO₂; and R^b is selected from hydrogen, carbocyclic and heterocyclic groups having from 3 to 12 ring members, and a C₁₋₈ hydrocarbyl group optionally substituted by one or more substituents selected from hydroxy, oxo, halogen, cyano, nitro, carboxy, amino, mono- or di-C₁₋₄ hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members and wherein one or more carbon atoms of the C₁₋₈ hydrocarbyl group may optionally be replaced by O, S, SO, SO₂, NR^c, X¹C(X²), C(X²)X¹ or X¹C(X²)X¹; or two adjacent groups R¹⁰, together with the carbon atoms or heteroatoms to which they are attached may form a 5-membered heteroaryl ring or a 5- or 6-membered non-aromatic heterocyclic ring, wherein the said heteroaryl and heterocyclic groups contain up to 3 heteroatom ring members selected from N, O and S;

R^c is selected from hydrogen and C₁₋₄ hydrocarbyl; and

X¹ is O, S or NR^c and X² is =O, =S or =NR^c;

R⁴ and R⁵ are the same or different and are selected from hydrogen and methyl; or one of R⁴ and R⁵ is selected from hydroxymethyl and ethyl and the other is hydrogen; and

R⁶ and R⁷ are the same or different and are selected from hydrogen and methyl;

together with a pharmaceutically acceptable carrier.

95. (new) A method for the prophylaxis or treatment of a disease state or condition mediated by a p38 MAP kinase, wherein the disease state or condition mediated by a p38 MAP kinase is selected from:

- (i) inflammatory and arthritic diseases and conditions, Reiter's syndrome, acute synovitis, rheumatoid arthritis, osteoarthritis, rheumatoid spondylitis, gouty arthritis, traumatic arthritis, rubella arthritis, psoriatic arthritis, graft vs. host reaction and allograft rejections;
- (ii) chronic inflammatory lung diseases, emphysema, chronic pulmonary inflammatory disease, chronic obstructive pulmonary disease (COPD), adult

respiratory distress syndrome and acute respiratory distress syndrome (ARDS);

(iii) lung diseases and conditions, tuberculosis, silicosis, pulmonary sarcoidosis, pulmonary fibrosis and bacterial pneumonia;

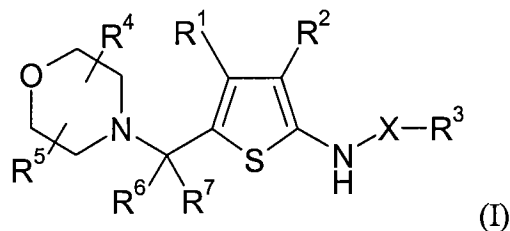
(iv) inflammatory diseases and conditions of the enteric tract, inflammatory bowel disease, Crohn's disease and ulcerative colitis;

(v) toxic shock syndrome and related diseases and conditions, sepsis, septic shock, endotoxic shock, gram negative sepsis and the inflammatory reaction induced by endotoxin;

(vi) Alzheimer's disease;

(vii) reperfusion injury;

(vii) diseases and conditions selected from atherosclerosis; muscle degeneration; gout; cerebral malaria; bone resorption diseases; fever and myalgias due to infection, influenza; cachexia, cachexia secondary to infection or malignancy, cachexia secondary to acquired immune deficiency syndrome (AIDS); AIDS; ARC (AIDS related complex); keloid formation; scar tissue formation; pyresis and asthma; which method comprises administering to a subject in need thereof a compound of the formula (I):



or a salt, solvate or N-oxide thereof, wherein:

R^1 and R^2 are the same or different and each is selected from hydrogen, saturated C_{1-3} hydrocarbyl, halogen and cyano;

X is selected from $C=O$, $C=S$, $C(=O)NH$, $C(=S)NH$, $C(=O)O$, $C(=O)S$, $C(=S)O$ and $C(=S)S$;

R^3 is selected from aryl and heteroaryl groups each having from 5 to 12 ring members and being unsubstituted or substituted by one or more substituent groups R^{10} ;

R^{10} is selected from halogen, hydroxy, trifluoromethyl, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members; a group R^a-R^b wherein R^a is a bond, O, CO, $X^1C(X^2)$, $C(X^2)X^1$, $X^1C(X^2)X^1$, S, SO, SO_2 , NR^c , SO_2NR^c or NR^cSO_2 ; and R^b is selected from hydrogen, carbocyclic and heterocyclic groups having from 3 to 12 ring members, and a C_{1-8} hydrocarbyl group optionally substituted by one or more substituents selected from hydroxy, oxo, halogen, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members and wherein one or more carbon atoms of the C_{1-8} hydrocarbyl group may optionally be replaced by O, S, SO, SO_2 , NR^c , $X^1C(X^2)$, $C(X^2)X^1$ or $X^1C(X^2)X^1$; or two adjacent groups R^{10} , together with the carbon atoms or heteroatoms to which they are attached may form a 5-membered heteroaryl ring or a 5- or 6-membered non-aromatic heterocyclic ring, wherein the said heteroaryl and heterocyclic groups contain up to 3 heteroatom ring members selected from N, O and S;

R^c is selected from hydrogen and C_{1-4} hydrocarbyl; and

X^1 is O, S or NR^c and X^2 is =O, =S or = NR^c ;

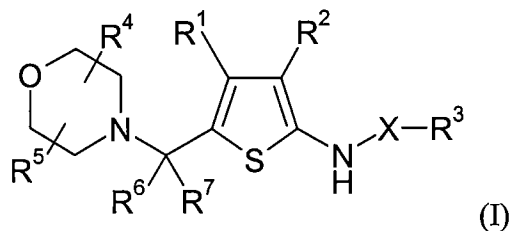
R^4 and R^5 are the same or different and are selected from hydrogen and methyl; or one of R^4 and R^5 is selected from hydroxymethyl and ethyl and the other is hydrogen; and

R^6 and R^7 are the same or different and are selected from hydrogen and methyl.

96. (new) A method according to claim 95 wherein the disease state or condition is selected from inflammatory diseases and conditions, rheumatoid arthritis and osteoarthritis.

97. (new) A method according to claim 95 wherein the disease state or condition is chronic obstructive pulmonary disease (COPD).

98. (new) A method for treating a disease or condition comprising or arising from abnormal cell growth in a mammal, the method comprising administering to the mammal a therapeutically effective amount of a compound of the formula (I):



or a salt, solvate or N-oxide thereof, wherein:

R^1 and R^2 are the same or different and each is selected from hydrogen, saturated C_{1-3} hydrocarbyl, halogen and cyano;

X is selected from C=O, C=S, C(=O)NH, C(=S)NH, C(=O)O, C(=O)S, C(=S)O and C(=S)S;

R^3 is selected from aryl and heteroaryl groups each having from 5 to 12 ring members and being unsubstituted or substituted by one or more substituent groups R^{10} ;

R^{10} is selected from halogen, hydroxy, trifluoromethyl, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members; a group R^a-R^b wherein R^a is a bond, O, CO, $X^1C(X^2)$, $C(X^2)X^1$, $X^1C(X^2)X^1$, S, SO, SO_2 , NR^c , SO_2NR^c or NR^cSO_2 ; and R^b is selected from hydrogen, carbocyclic and heterocyclic groups having from 3 to 12 ring members, and a C_{1-8} hydrocarbyl group optionally substituted by one or more substituents selected from hydroxy, oxo, halogen, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members and wherein one or more carbon atoms of the C_{1-8} hydrocarbyl group may optionally be replaced by O, S, SO, SO_2 , NR^c , $X^1C(X^2)$, $C(X^2)X^1$ or $X^1C(X^2)X^1$; or two adjacent groups R^{10} , together with the carbon atoms or heteroatoms to which they are attached may form a 5-membered heteroaryl ring or a 5- or 6-membered non-aromatic heterocyclic ring, wherein the said heteroaryl and heterocyclic groups contain up to 3 heteroatom ring members selected from N, O and S;

R^c is selected from hydrogen and C_{1-4} hydrocarbyl; and

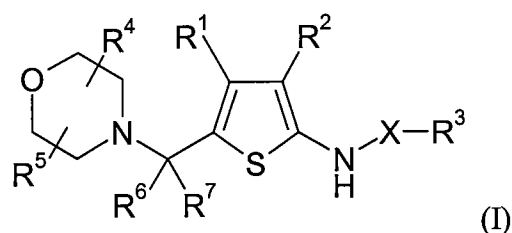
X^1 is O, S or NR^c and X^2 is =O, =S or = NR^c ;

R^4 and R^5 are the same or different and are selected from hydrogen and methyl; or one of R^4 and R^5 is selected from hydroxymethyl and ethyl and the other is

hydrogen; and

R^6 and R^7 are the same or different and are selected from hydrogen and methyl.

99. (new) A method for the prophylaxis or treatment of a disease state or condition mediated by a raf kinase, which method comprises administering to a subject in need thereof a compound of the formula (I):



or a salt, solvate or N-oxide thereof, wherein:

R^1 and R^2 are the same or different and each is selected from hydrogen, saturated C_{1-3} hydrocarbyl, halogen and cyano;

X is selected from C=O, C=S, C(=O)NH, C(=S)NH, C(=O)O, C(=O)S, C(=S)O and C(=S)S;

R^3 is selected from aryl and heteroaryl groups each having from 5 to 12 ring members and being unsubstituted or substituted by one or more substituent groups R^{10} ;

R^{10} is selected from halogen, hydroxy, trifluoromethyl, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members; a group R^a-R^b wherein R^a is a bond, O, CO, $X^1C(X^2)$, $C(X^2)X^1$, $X^1C(X^2)X^1$, S, SO, SO_2 , NR^c , SO_2NR^c or NR^cSO_2 ; and R^b is selected from hydrogen, carbocyclic and heterocyclic groups having from 3 to 12 ring members, and a C_{1-8} hydrocarbyl group optionally substituted by one or more substituents selected from hydroxy, oxo, halogen, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members and wherein one or more carbon atoms of the C_{1-8} hydrocarbyl group may optionally be replaced by O, S, SO, SO_2 , NR^c , $X^1C(X^2)$, $C(X^2)X^1$ or $X^1C(X^2)X^1$; or two adjacent groups R^{10} , together with the carbon atoms or heteroatoms to which

they are attached may form a 5-membered heteroaryl ring or a 5- or 6-membered non-aromatic heterocyclic ring, wherein the said heteroaryl and heterocyclic groups contain up to 3 heteroatom ring members selected from N, O and S;

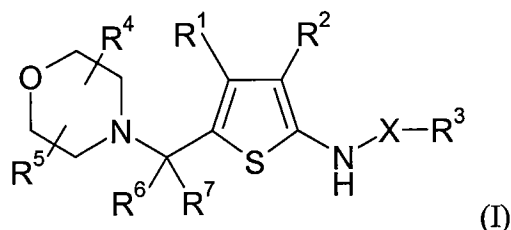
R^c is selected from hydrogen and C_{1-4} hydrocarbyl; and

X^1 is O, S or NR^c and X^2 is $=O$, $=S$ or $=NR^c$;

R^4 and R^5 are the same or different and are selected from hydrogen and methyl; or one of R^4 and R^5 is selected from hydroxymethyl and ethyl and the other is hydrogen; and

R^6 and R^7 are the same or different and are selected from hydrogen and methyl.

100. (new) A process for the preparation of a compound of the formula (I):



or a salt, solvate or N-oxide thereof, wherein:

R^1 and R^2 are the same or different and each is selected from hydrogen, saturated C_{1-3} hydrocarbyl, halogen and cyano;

X is selected from $C=O$, $C=S$, $C(=O)NH$, $C(=S)NH$, $C(=O)O$, $C(=O)S$, $C(=S)O$ and $C(=S)S$;

R^3 is selected from aryl and heteroaryl groups each having from 5 to 12 ring members and being unsubstituted or substituted by one or more substituent groups R^{10} ;

R^{10} is selected from halogen, hydroxy, trifluoromethyl, cyano, nitro, carboxy, amino, mono- or di- C_{1-4} hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members; a group R^a-R^b wherein R^a is a bond, O, CO, $X^1C(X^2)$, $C(X^2)X^1$, $X^1C(X^2)X^1$, S, SO, SO_2 , NR^c , SO_2NR^c or NR^cSO_2 ; and R^b is selected from hydrogen, carbocyclic and heterocyclic groups having from 3 to 12 ring members, and a C_{1-8} hydrocarbyl group optionally substituted by one or more

substituents selected from hydroxy, oxo, halogen, cyano, nitro, carboxy, amino, mono- or di-C₁₋₄ hydrocarbylamino, carbocyclic and heterocyclic groups having from 3 to 12 ring members and wherein one or more carbon atoms of the C₁₋₈ hydrocarbyl group may optionally be replaced by O, S, SO, SO₂, NR^c, X¹C(X²), C(X²)X¹ or X¹C(X²)X¹; or two adjacent groups R¹⁰, together with the carbon atoms or heteroatoms to which they are attached may form a 5-membered heteroaryl ring or a 5- or 6-membered non-aromatic heterocyclic ring, wherein the said heteroaryl and heterocyclic groups contain up to 3 heteroatom ring members selected from N, O and S;

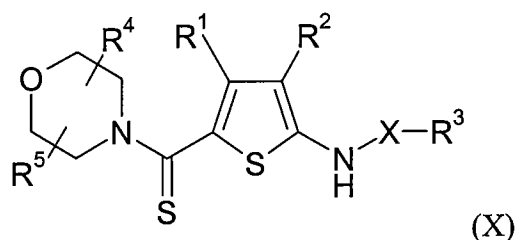
R^c is selected from hydrogen and C₁₋₄ hydrocarbyl; and

X¹ is O, S or NR^c and X² is =O, =S or =NR^c;

R⁴ and R⁵ are the same or different and are selected from hydrogen and methyl; or one of R⁴ and R⁵ is selected from hydroxymethyl and ethyl and the other is hydrogen; and

R⁶ and R⁷ are the same or different and are selected from hydrogen and methyl;

which process comprises the S-alkylation of a compound of the formula (X):



using an alkylating agent to give a thioimide intermediate followed by:

(i) reduction of the thioimide intermediate to give a compound of formula (I) in which R⁶ and R⁷ are hydrogen by means of a reducing agent; or

(ii) treating the thioimide intermediate with methyl lithium or a methyl Grignard reagent, followed by a reducing agent to give a compound of the formula (I) in which one of R⁶ and R⁷ is methyl; or

(iii) treating the thioimide intermediate with more than one equivalent of methyl lithium or a methyl Grignard reagent to give a compound of the formula (I) in which both R⁶ and R⁷ are methyl.